#### XII. APPENDIX IV

#### MATERIAL SAFETY DATA SHEET

The following items of information which are applicable to a specific product or material shall be provided in the appropriate block of the Material Safety Data Sheet (MSDS).

The product designation is inserted in the block in the upper left corner of the first page to facilitate filing and retrieval. Print in upper case letters as large as possible. It should be printed to read upright with the sheet turned sideways. The product designation is that name or code designation which appears on the label, or by which the product is sold or known by employees. The relative numerical hazard ratings and key statements are those determined by the rules in Chapter V, Part B, of the NIOSH publication, An Identification System for Occupationally Hazardous Materials. The company identification may be printed in the upper right corner if desired.

#### (a) Section I. Product Identification

The manufacturer's name, address, and regular and emergency telephone numbers (including area code) are inserted in the appropriate blocks of Section I. The company listed should be a source of detailed backup information on the hazards of the material(s) covered by the MSDS. The listing of suppliers or wholesale distributors is discouraged. The trade name should be the product designation or common name associated with the material. The synonyms are those commonly used for the product, especially formal chemical nomenclature. Every known chemical designation or

competitor's trade name need not be listed.

## (b) Section II. Hazardous Ingredients

The "materials" listed in Section II shall be those substances which are part of the hazardous product covered by the MSDS and individually meet any of the criteria defining a hazardous material. Thus, one component of a multicomponent product might be listed because of its toxicity, another component because of its flammability, while a third component could be included both for its toxicity and its reactivity. Note that a MSDS for a single component product must have the name of the material repeated in this section to avoid giving the impression that there are no hazardous ingredients.

Chemical substances should be listed according to their complete name derived from a recognized system of nomenclature. Where possible, avoid using common names and general class names such as "aromatic amine," "safety solvent," or "aliphatic hydrocarbon" when the specific name is known.

The "%" may be the approximate percentage by weight or volume (indicate basis) which each hazardous ingredient of the mixture bears to the whole mixture. This may be indicated as a range or maximum amount, ie, "10-40% vol" or "10% max wt" to avoid disclosure of trade secrets.

Toxic hazard data shall be stated in terms of concentration, mode of exposure or test, and animal used, eg, "100 ppm LC50-rat," "25 mg/kg LD50-skin-rabbit," "75 ppm LC man," or "permissible exposure from 29 CFR 1910.1000," or if not available, from other sources of publications such as the American Conference of Governmental Industrial Hygienists or the American National Standards Institute Inc. Flashpoint, shock sensitivity

or similar descriptive data may be used to indicate flammability, reactivity, or similar hazardous properties of the material.

#### (c) Section III. Physical Data

The data in Section III should be for the total mixture and should include the boiling point and melting point in degrees Fahrenheit (Celsius in parentheses); vapor pressure, in conventional millimeters of mercury (mmHg); vapor density of gas or vapor (air = 1); solubility in water, in parts/hundred parts of water by weight; specific gravity (water = 1); percent volatiles (indicated if by weight or volume) at 70 degrees Fahrenheit (21.1 degrees Celsius); evaporation rate for liquids or sublimable solids, relative to butyl acetate; and appearance and odor. These data are useful for the control of toxic substances. Boiling point, vapor density, percent volatiles, vapor pressure, and evaporation are useful for designing proper ventilation equipment. This information is also useful for design and deployment of adequate fire and containment equipment. The appearance and odor may facilitate identification of substances stored in improperly marked containers, or when spilled.

#### (d) Section IV. Fire and Explosion Data

Section IV should contain complete fire and explosion data for the product, including flashpoint and autoignition temperature in degrees Fahrenheit (Celsius in parentheses); flammable limits, in percent by volume in air; suitable extinguishing media or materials; special firefighting procedures; and unusual fire and explosion hazard information. If the product presents no fire hazard, insert "NO FIRE HAZARD" on the line labeled "Extinguishing Media."

#### (e) Section V. Health Hazard Information

The "Health Hazard Data" should be a combined estimate of the hazard of the total product. This can be expressed as a TWA concentration, as a permissible exposure, or by some other indication of an acceptable standard. Other data are acceptable, such as lowest LD50 if multiple components are involved.

Under "Routes of Exposure," comments in each category should reflect the potential hazard from absorption by the route in question. Comments should indicate the severity of the effect and the basis for the statement if possible. The basis might be animal studies, analogy with similar products, or human experiences. Comments such as "yes" or "possible" are not helpful. Typical comments might be:

Skin Contact--single short contact, no adverse effects likely; prolonged or repeated contact, possibly mild irritation.

Eye Contact--some pain and mild transient irritation; no corneal scarring.

"Emergency and First Aid Procedures" should be written in lay language and should primarily represent first-aid treatment that could be provided by paramedical personnel or individuals trained in first aid.

Information in the "Notes to Physician" section should include any special medical information which would be of assistance to an attending physician including required or recommended preplacement and periodic medical examinations, diagnostic procedures, and medical management of overexposed employees.

#### (f) Section VI. Reactivity Data

The comments in Section VI relate to safe storage and handling of hazardous, unstable substances. It is particularly important to highlight instability or incompatibility to common substances or circumstances, such as water, direct sunlight, steel or copper piping, acids, alkalies, etc. "Hazardous Decomposition Products" shall include those products released under fire conditions. It must also include dangerous products produced by aging, such as peroxides in the case of some ethers. Where applicable, shelf life should also be indicated.

#### (g) Section VII. Spill or Leak Procedures

Detailed procedures for cleanup and disposal should be listed with emphasis on precautions to be taken to protect employees assigned to cleanup detail. Specific neutralizing chemicals or procedures should be described in detail. Disposal methods should be explicit including proper labeling of containers holding residues and ultimate disposal methods such as "sanitary landfill" or "incineration." Warnings such as "comply with local, state, and federal antipollution ordinances" are proper but not sufficient. Specific procedures shall be identified.

#### (h) Section VIII. Special Protection Information

Section VIII requires specific information. Statements such as "Yes," "No," or "If necessary" are not informative. Ventilation requirements should be specific as to type and preferred methods. Respirators shall be specified as to type and NIOSH or US Bureau of Mines approval class, ie, "Supplied air," "Organic vapor canister," etc. Protective equipment must be specified as to type and materials of construction.

#### (i) Section IX. Special Precautions

"Precautionary Statements" shall consist of the label statements selected for use on the container or placard. Additional information on any aspect of safety or health not covered in other sections should be inserted in Section IX. The lower block can contain references to published guides or in-house procedures for handling and storage. Department of Transportation markings and classifications and other freight, handling, or storage requirements and environmental controls can be noted.

#### (j) Signature and Filing

Finally, the name and address of the responsible person who completed the MSDS and the date of completion are entered. This will facilitate correction of errors and identify a source of additional information.

The MSDS shall be filed in a location readily accessible to employees exposed to the hazardous material. The MSDS can be used as a training aid and basis for discussion during safety meetings and training of new employees. It should assist management by directing attention to the need for specific control engineering, work practices, and protective measures to ensure safe handling and use of the material. It will aid the safety and health staff in planning a safe and healthful work environment and in suggesting appropriate emergency procedures and sources of help in the event of harmful exposure of employees.

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MATERIAI	SAFE	TY D	ATA	SHEET
I PROD	UCT IDENTI	FICATIO	N	
MANUFACTURER'S NAME			TELEPHONE N	
ADDRESS				
TRADE NAME				
SYNONYMS				
II HAZA	RDOUS ING	REDIEN	rs	
MATERIAL OR COMPON	IENT		*	HAZARD DATA
		······		
	<u>.</u> ,			
111	PHYSICAL I	DATA		
BOILING POINT, 760 MM HG		MELTING PO	INT	
SPECIFIC GRAVITY (H20=1)		VAPOR PRE	SSURE	
VAPOR DENSITY (AIR=1)		SOLUBILITY	' IN H2O, % BY	/ WT
% VOLATILES BY VOL		EVAPORATI	ON RATE (BU	TYL ACETATE=1}
APPEARANCE AND ODOR				

	IV FIRE AND	EXPLO	SION DATA	
FLASH POINT (TEST METHOD)			AUTOIGNITION TEMPERATURE	
FLAMMABLE LIMITS IN AI	R, % BY VOL.	LOWER		UPPER
EXTINGUISHING MEDIA				
SPECIAL FIRE FIGHTING PROCEDURES				
UNUSUAL FIRE AND EXPLOSION HAZARD				
	V HEALTH HA	ZARD II	FORMATIO	N
HEALTH HAZARD DATA	V			
ROUTES OF EXPOSURE				
INHALATION				
SKIN CONTACT				
SKIN ABSORPTION				
EYE CONTACT			····	
INGESTION				
EFFECTS OF OVEREXPOSU ACUTE OVEREXPOS				
CHRONIC OVEREXP	OSURE			
EMERGENCY AND FIRST A	ND PROCEDURES			
EYES				
SKIN:				
INHALATION:				
INGESTION	· · · · · · · · · · · · · · · · · · ·		· · · · · · · · · · · · · · · · · · ·	
NOTES TO PHYSICIAN				

VI REACTIVITY DATA
CONDITIONS CONTRIBUTING TO INSTABILITY
INCOMPATIBILITY
HAZARDOUS DECOMPOSITION PRODUCTS
CONDITIONS CONTRIBUTING TO HAZARDOUS POLYMERIZATION
VII SPILL OR LEAK PROCEDURES
STEPS TO BE TAKEN IF MATERIAL IS RELEASED OR SPILLED
NEUTRALIZING CHEMICALS
WASTE DISPOSAL METHOD
VIII SPECIAL PROTECTION INFORMATION
VENTILATION REQUIREMENTS
SPECIFIC PERSONAL PROTECTIVE EQUIPMENT
RESPIRATORY (SPECIFY IN DETAIL)
EYE
GLOVES
OTHER CLOTHING AND EQUIPMENT

	IX SPECIAL PRECAUTIONS	
PRECAUTIONARY STATEMENTS		
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OTHER HANDLING AND STORAGE REQUIREMENTS		
PREPARED BY		
ADDRESS:		
DATE.		

#### XIII. TABLES AND FIGURE

#### TABLE XIII-1

#### PHYSICAL PROPERTIES OF CARBARYL

Appearance	White, crystalline solid
Empirical formula	C12H11O2N
Formula weight	201.22
Flashpoint, open cup (Cleveland)	193 C
Melting point	142 C
Specific gravity	1.232 at 20 C
Vapor pressure	0.000041 mmHg at 25 C, 0.00015 mmHg at 40 C
Explosibility (minimum concentration of technical grade dust)	0.02 oz/cu ft (20.3 g/cu m)
Solubility in water	40 ppm at 30 C
Solubility in organic solvents	Moderately soluble in acetone, N,N-dimethyl formamide, isophorone and cyclohexanone
Stability	Stable to light, heat, and acids; hydrolyzed in alkalies

Adapted from Kirk-Othmer, [11] Merck Index, [12] Union Carbide Corporation, [13 (sec 16)] and HH Moorefield (written communication, February 1976)

#### TABLE XIII-2

#### SYNONYMS AND TRADE NAMES FOR CARBARYL

Atoxan Caprolin Carpolin Compound 7744 Crag Sevin Experimental insecticide 7744 Gamonil N-methyl-alpha-naphthylcarbamate N-methyl-alpha-naphthylurethan N-methyl-1-napthyl carbamate 1-Naphthol N-methylcarbamate Alpha-naftyl-N-methylkarbamat (Czech) Alpha-naphthyl N-methylcarbamate 1-Naphthyl N-methylcarbamate Panam Sevidol Sevin Union Carbide 7,744

Compiled from the Registry of Toxic Effects of Chemical Substances, 1975 edition [14]

CONCENTRATIONS (MG/KG) OF CARBARYL
IN RABBIT TISSUES AND EXCRETORY FLUIDS AFTER
SINGLE ORAL DOSES OF 400 MG/KG OF CARBARYL

TABLE XIII-3

Sample	Hours after Administration							
	0.5	1	2.5	4	18	24	48	
Brain	0.0075	0	0.03	0.054	0	0.215	0.03	
Medulla oblongata	0.015	_	0.03	0.09	-	-	0	
Cerebellum	0.021	_	0.03	_	_	_	0	
Kidneys	0.03	0.75	0.75	0.18	1.125	0	0	
Spleen	0.075	0.075	_	0.036	0	0	0	
Liver	0.225	0.975	0.15	0.036	0.15	0	0.075	
Heart	0.3	0	0.03	0.018	0	0	0	
Lungs	0.003	0	0	0.015	0	0	0	
Lumbar muscles	0.03	0.03	0	0.012	0	0	0	
Femoral muscles	0.03	0	0	0	0	0	0	
Bile	3.75	0.075	4.5	0.18	1.5	1.5	1.5	
Urine	1.5	1.125	11.25	1.8	0	0.75	0	
Testes	0.075	0	0	0.09	_	0	0	
Kidney fat	1.25	_	0.6	0.036	-	0	0.03	

Adapted from Bukin and Filatov [86]

TABLE XIII-4

CARBARYL METABOLITES FROM MAMMALIAN\* URINE

Metabolites		Labeled Forms of Carbary1**						Unlabeled Carbaryl***								
	1		140 ph	C- thy	71				140 et1	C- hy:	L		14C- Carbony1			
Unidentified neutrals	R	G	M	P	s	D	R	G	М	P	s	D	R	R	Но	Hi
l-Naphthyl methylcarbamate N-glucuronide		G						G						R		
Unidentified metabolite (A)	R	G				D	R	G	M			D	R	R	Но	
l-Naphthyl methylimido- carbonate-o-glucuronide	R	G	M	P	S	D	R	G	M	P	s	D	R	R	Но	
Unidentified metabolite (B)		G				D		G				D				
4-(Methylcarbamoyloxy)- l-naphthyl glucuronide	R	G	M	P	s		R	G	M	P	s		R	R	Но	
1-Naphthyl glucuronide	R	G		P	S									R	Но	H1
4-(Methylcarbamoyloxy)- l-naphthyl sulfate	R	G	M		S		R	G	M	I	s		R	R		
1-Naphthyl sulfate	R	G	M		S									R	Но	Hi
Unidentified metabolite (C)											S					
Unidentified metabolite (D)					S						S					

<sup>\*</sup>Rat (R), guinea pig (G), monkey (M), swine (P), sheep (S), dog (D), and human (H)

\dapted from Knaak et al [33,34] and Knaak and Sullivan [87]

<sup>\*\*</sup>Analysis by radiometric techniques; rats and guinea pigs treated ip, all others orally

<sup>\*\*\*</sup>Analysis by fluorometric techniques; rats treated orally, humans orally (o) and by inhalation (i)

EFFECTS OF CARBARYL EXPOSURE ON HUMANS

TABLE XIII-5

Routes of Exposure	Subjects	Exposure Concentrat and Duration	ion Effects	Refer- ence
Respiratory and possi- bly dermal	59 workers	0.03 - 40 mg/cu m 16 mo (hourly expo- sure not stated) 8 mo (hourly expo- sure not stated)	Whole blood ChE* activities inhibited but within normal range; urinary 1-naphthol 1,000 µg/100 ml** or higher in 41% of samples (conc low on Mon, rising during workweek, in substudy of 7 workers)	28 t
11	Agricul- tural workers (number unknown)	2 - 4 mg/cu m 4 - 6 hr/d, 3 - 4 d	Whole blood ChE activities inhibited 11-30%; other clinical tests negative	39
11	10 sprayers	Insecticide (unknown conc) single application	Rash in 1 splashed worker; plasma ChE slightly inhibited on d 1, normal on d 5 after application in all 10	35
**	95 exposed villagers	Insecticide (unknown conc) 1 wk	Plasma ChE activity still inhibited l wk after spraying	35
11	l man (with res- pirator)	40.6 - 49.3 mg/cu m for 2 d	Urinary 1-naphthol 2,340 and 8,975 µg/100 ml	13 (sec 9, 10)
**	l man (no res- pirator)	45.2 - 50.9 mg/cu m for 2 d	Urinary 1-naphthol 2,340 and 3,619 µg/100 ml	13 (sec 9, 10)
n	1 man	Carbaryl-sulfur mixture (unknown conc and duration)	Weakness, dizziness, shortness of breath on day of exposure and 2 d later	38

Routes of Exposure	Subjects	Exposure Concentrat and Duration	ion Effects	Refer- ence
Respiratory and possi- bly dermal	14 exposed male workers	Dust (unknown conc) 8 - 16 hr	Urinary 1-naphthol 2,443 µg/100 ml for 7 asymptomatic workers; 1,417 for 7 symptomatic workers, all with dizziness, nausea, 1 with headache, 1 with overheating, perspiration	13 (sec 7)
11	Workers (number unknown)	Dust (unknown conc)	Urinary 1-naphthyl glucuronide and 1-naphthyl sulfate higher during exposure than 72 hr after exposure	34
Dermal	6 men	4 μg/sq cm 14C-carbaryl in acetone to forearm	Recovery in urine, 73.9% of dose over 5 d	30
11	6 men	4 μg/sq cm 14C-carbaryl in acetone to angle of jaw	Recovery in urine, 70% of dose over 5 d	31
Oral	2 men	2 mg/kg	Recovery in urine, 26-27% of dose in 24 hr as metabolites, 1-naphthyl sulfate, 1-naphthyl glucuronid 4-(methylcarbamoyloxy -1 naphthyl glucuroni	e, ')
"	2 men 2 men 2 men	0.5 mg/kg 1.0 mg/kg 2.0 mg/kg (single doses)	No change in plasma or whole blood ChE; no toxic symptoms	32

Routes of Exposure	Subjects	Exposure Concentrate and Duration	ion Effects	Refer- ence
Oral	5 men	0.06 mg/kg/d 6 wk	Urinary rise of aa/cr***	32
11	5 men	0.12 mg/kg/d 6 wk	Urinary rise of aa/cr***	32
"	l man	2.8 mg/kg (form unstated)	Epigastric pain, sweating; recovery in 2 hr with atro- pine therapy	21
"	l child (19 mo old)	Solution (unknown conc)	Miosis, salivation, incoordination; recovery in 12 hr with atropine therapy	28
11	1 man	Solution (unknown conc; ate watermelon sprayed with 80% carbaryl)	Nausea, vomiting hyperreflexia, pallor intestinal colic, na- sal discharge; recov- ery in 18 hr with deoxycorticosterone	<b>41</b>
**	1 man	80% solution (several ml)	Nausea, salivation, headache, tremors, lacrimation; recov- ery in 1.5 hr with deoxycorticosterone	41
***	1 man	80% solution 0.5 liter	Whole blood ChE strongly inhibited; death in 6 hr; carbaryl present in gastrointestinal trace blood, liver, kid- neys, and urine	40 t,

<sup>\*</sup>Cholinesterase

<sup>\*\*</sup>Normal range, 150-400  $\mu$ g/100 ml urine \*\*\*Amino acid nitrogen to creatinine ratio

TABLE XIII-6

EFFECTS OF CARBARYL EXPOSURE ON ANIMALS

Routes of Exposure		No.	Exposure Concentratio	Effects	Refer- ence
Respira- tory	Guinea pigs	6	390 mg/cu m (dust) 4 hr	Nasal and eye irrita- tion; hemorrhagic areas in lungs	27
"	ff	6	230 mg/cu m (dust) 4 hr	Slight initial weight loss; recovery by d 14	27
11	Rats	-	10 mg/cu m (dust) 7 hr/d 5 d/wk for 90 d	No effects	27
**	Dogs	-	75 mg/cu m (dust) (duration unknown)	Typical signs of ChE** inhibition within 5 hr	
**	Cats	4	82 mg/cu m (dust) 6 hr	Signs of toxicity observed; ChE in- hibited, normal after 72 hr	39
**	**	4	37 mg/cu m (dust) 6 hr	ChE inhibited, normal after 48 hr	39
II .	**	4	20 mg/cu m (dust) 6 hr	ChE inhibited, normal after 24 hr	39
11	11	4	63 mg/cu m (dust) 6 hr/d for 1 mo	Salivation; ChE in- hibited 31-40% (ser- um), 41-58% (erythro- cyte); 1 death at d 20	39
11	***	4	40 mg/cu m (dust) 6 hr/d for 2 mo	Up to 50% erythrocyte ChE inhibition; de- terioration of condi- tioned reflexes (un- defined)	39
11	11	4	<pre>16 mg/cu m (dust) 6 hr/d for 4 mo</pre>	No effects	39

Routes of Exposure	Species	No.	Exposure Concentration and Duration*	n Effects	Refer- ence
Ocular	Rabbits	5	0.5 ml of 10% suspension applied to 1 eye of each animal	Mild irritation in 1 eye after 24 hr	27
11	**	5	50 mg dust	Traces of corneal necrosis	27
11	77	5	25% suspension in water	No injury	27
11	**	-	10% suspension in water, or 50 mg powder	Transient miosis, hyperemia	39
Derma1	11	5	0.01 ml of 10% soln in acetone	No irritation	27
11	**	6	500 mg/kg (form unspecified)	Inhibition of serum and erythrocyte ChE in 1st 24 hr	39
II	Rats	10 M 10 F	Various doses in xylene	LD50 greater than 4,000 mg/kg in both sexes	49
Intracu- taneous	Guinea pigs	16	0.1 ml (0.1% in propylene glycol) 8 alternate days (3 wk after last dose, another challenge dose injected)	Weak sensitization in 4 animals	27
Subcuta- neous	Rats	5-9 per group	10 mg/kg/wk 2 - 5 wk	Behavioral changes; decreased motivation and response to elec- troshock	57

Routes o		No.	Exposure Concentration and Duration*	n <b>Effects</b>	Refer- ence
Subcuta- neous	Mice	60 M	10 mg/mouse/wk in 0.25% agar 5 mo	No increases in tu- mors, infections, or mortality compared to control animals	27
11	Chickens	13	0.25 - 3 g/kg (suspended in lard)	Leg weakness observed at 2 g/kg; no effects at doses of 1 g/kg or less	
***	Chickens (atropin- ized)	<del></del>	0.8 - 1.6 g/kg	Leg weakness; recovery by d 24	43
Íр	Rats	-	0.56 mg/kg and 2.24 mg/kg	Decreased physical activity; reversed with atropine therapy	58
iv	Dogs	6	<pre>10 and 15 mg/kg in ethyl alcohol</pre>	No change in erythrocyte or plasma ChE activity	27
***	ti	5	30 mg/kg	Lacrimation, saliva- tion, tremors	55
"	Miniature pigs	8 M	20 mg/kg	Tremors, ataxia, in- coordination, para- plegia	55
Oral	Guinea pigs (preg- nant)	26 F	300 mg/kg/d in gelatin capsule d ll - 20 of gestation	Maternal mortality 38%; fetal mortality 17.5% in survivors' litters (9.5% in con- trols); ll terata in fetuses of treated	71

Routes of Exposure		No.	Exposure Concentration and Duration*	n Effects	Refer- ence
Oral	Guinea pigs (preg- nant)	40 F	300 mg/kg in capsule single dose d 11 - 20 of gestation	Maternal mortality 12.5%; fetal mortal- ity 6.5% (controls 9.5%); 9 malformed fetuses in litters of treated	71
***	11	150 F	100, 200, 300 mg/kg in diet 1- to 15-d intervals during d 10 - 24 of gestation	Weight gains less than controls; no significant increase in terata compared to controls	72
11	11	150 F	50, 100, 200 mg/kg by intubation 1- to 15-d intervals during d 10 - 24 of gestation	Increased fetal anom- alies at 50 mg/kg only (15.7% vs 9.1% in controls)	72
***	Rabbits (preg- nant)	17 F	50, 100, 200 mg/kg in capsules on d 5 - 15 of gestation	No terata or dose- related mortality compared to controls	71
11	Rats (preg- nant)	18 F	500 mg/kg/d in diet during various intervals of gestation or until weaning of pups	Weight gain of off- spring reduced; 2/3 mortality of pups within 4 d after birt	70 h
***	***	36 F	20 and 100 mg/kg/d in diet during various intervals of gestation or until weaning of pups	No effect on viabil- ity of pups compared to controls	70

Routes of Exposure		No.	Exposure Concentratio and Duration*	n Effects	Refer- ence
Oral	Rats	- F	610 mg/kg (gavage)	LD50	27
11	***	-	510 mg/kg (gavage)	LD50	27
***	11	- M	850 mg/kg (gavage)	LD50	43
11	11	- M	600 mg/kg (gavage)	Lowest lethal dose	43
**	**	- F	500 mg/kg (gavage)	LD50	43
11	**	- F	100 mg/kg (gavage)	Lowest lethal dose	43
11	"	10 M,	F 2,250 ppm in diet 96 d	Cloudy swelling of kidney tubules in 4	27
***	11	20 F	400 ppm in diet 2 yr	Cloudy swelling of hepatic cords; cloudy swelling of kidney tubules present after 1 yr; incidence not significant compared to controls after 2 yr	27
11	11	60 F 80 M	50, 100, 200 ppm in diet, 2 yr	Effects seen similar to controls	27
**	"	48 M 48 F	14 and 70 mg/kg/d 12 mo	Increased gonado- tropic hormone, ad- renal cortex activity growth inhibition; blood ChE inhibited after 3 mo; testicu- lar changes; estrus cycle prolonged	77 <b>;</b>

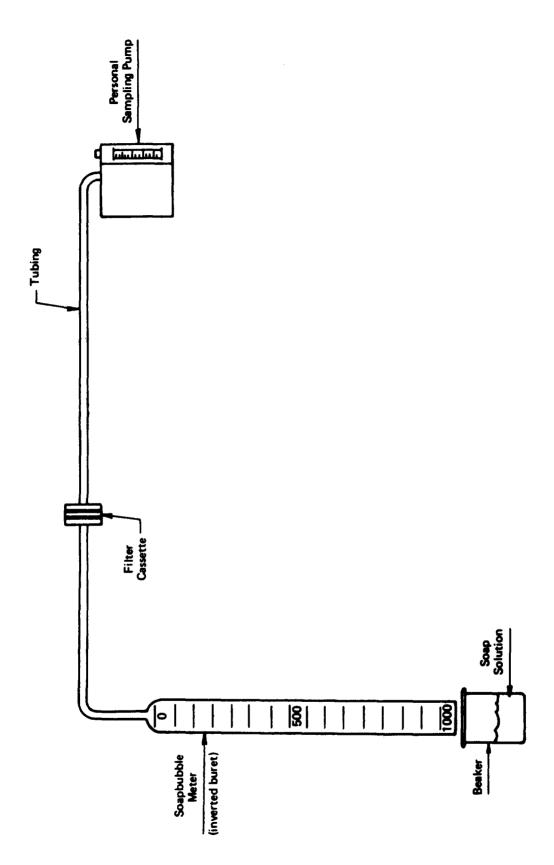
Routes of Exposure	Species	No.	Exposure Concentration and Duration*	n Effects	Refer- ence
Oral	Rats	24 M 24 F	7 mg/kg/d 12 mo	Increased gonado- tropic hormone, ad- renal cortex activity	77
"	Mice	20 M	50 - 1,000 mg/kg/d 5 d (gavage)	Death in 1 of 10 at 1,000 mg/kg; no evidence for dominant lethal mutagenicity after mating	59
**	Rats	60 (FO)	2,000, 5,000, 10,000 ppm in diet 3 generations	Weight below controls for all treated wean- lings; average litter size and survival of offspring decreased at 5,000- and 10,000- ppm levels	78
11	11	16 (F0)	200 mg/kg/d in diet 3 generations	Initial decrease in weight gain in FO; lengthened gestation period in F1 and F2 compared to controls	72
11	**	48 (F0)	7, 25, 100 mg/kg/d in diet 3 generations	No effects compared to controls in all generations	72
**	"	16 (F0)	100 mg/kg/d by gavage 3 generations	Increased fetal mor- tality; sporadic ad- verse effects on litte size, adult survival time, fetal resorption in all generations	
**	11	32 (F0)	3, 7, 25 mg/kg/d by gavage 3 generations	No effects compared to controls in all generations	72

Routes of Exposure		No.	Exposure Concentration and Duration*	n Effects	Refer- ence
Oral	Rats	- F	2 and 5 mg/kg/d in sunflower oil 4 generations (F2 to F5) 6 mo each	Fertility decreasing progressively, dose-related (F2 to F4); reduced estrus time in F3 and F4	73
11	11	- M	<pre>2 and 5   mg/kg/d in sunflower oil 3 generations  (F2 to F4) 6 mo each</pre>	Sperm motility, sper- matogenesis, duration of sperm survival reduced compared to controls	
***	Gerbils	108 (F0)	2,000 - 10,000 ppm in diet 3 generations	Decreased fertility, litter size, pup via- bility in all genera- tions; at and above 4,000 ppm, weanling weights less than controls	
***	Hamsters (preg- nant)	6 F	250 mg/kg/d (gavage) d 7 - 8 of gestation	Signs of ChE inhibi- tion; salivation, diarrhea, incoordina- tion; 2 deaths; fetal mortality 30.3% (con- trol 5.5%), no terato- logic defects in fetu	_
11	"	8 F	125 mg/kg/d (gavage) d 6 - 8 of gestation	Fetal mortality 10% (control 5.5%), no teratologic defects in fetuses	71

Routes of Exposure	=	No.	E	kposure Concentratio and Duration*	en Effects	Refer- ence
Oral	Mice (preg- nant)	20	F	30 mg/kg/d in diet from d 6 of gestation to delivery	Minor abnormalities in 9 fetuses (controls, 2)	75
"	11	20	F	10 mg/kg/d in diet from d 6 of gestation to delivery	No effects compared to controls	75
**	ti	36 36		4.64 mg/kg/d 18 mo (from d 7 of age)	No significant in- crease in tumors compared to controls	79
"	Dogs (preg- nant)	55	F	3.125 - 50 mg/kg/d (diet) throughout gestation	Dystocia in 1/3 of dams, none in control 11.6% terata at doses of 6.25 mg/kg or more	
11	Rhesus monkeys	10	F	20.0 mg/kg/d by gavage throughout gestation	Abortions 3 in 6 pregnancies (1 abortion in 5 pregnancies for 7 controls)	67
11	***	4	F	<pre>2.0 mg/kg/d by gavage   throughout   gestation</pre>	Abortions 2 in 2 pregnancies	67
11	11	16	đ	20 mg/kg/d in capsules 20 - 38 of gestation	Abortions 3 in 15 pregnancies, no terata	69
11	11	16	đ	2 mg/kg/d in capsules 20 - 38 of gestation	Abortions 1 in 16 pregnancies, no terata	69

Routes of Exposure		No.	Exposure Concentration and Duration*	n Effects	Refer- ence
Oral	Rhesus monkeys	16	0.2 mg/kg/d in capsules d 20 - 38 of gestation	Abortions 2 in 16 pregnancies, no terata (4 abortions, 1 stillborn in 31 pregnancies for 31 controls; no terata)	69
***************************************	Pigs	1 M 1 F	<b>5</b> . <b>5</b> .	First signs of toxi- city at d 45 and 62; ataxia, incoordination tremors, vascular degeneration, myopath cerebral edema; death on d 72 (M) and 83 (F	у,
11	"	2 M 1 F		First signs of toxicity at d 37, 39, 42; ataxia, incoordination, tremors; vascular degeneration, myopathy, cerebral edema; death on d 46 (2 M) and 85 (F)	52
11	Miniature pigs	e 5 M	125 mg/kg/d in diet 6 - 8 wk	Spastic paresis of posterior extremities	55

<sup>\*</sup>Single dose unless duration is specified \*\*Cholinesterase



# DEPARTMENT OF HEALTH, EDUCATION, AND WELFARE

PUBLIC HEALTH SERVICE

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